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<!--StartFragment-->RESULT 1
US-08-690-102A-4
; Sequence 4, Application US/08690102A
; Patent No. 5789554
; GENERAL INFORMATION:
; APPLICANT: LEUNG, Shui-on
; APPLICANT: HANSEN, Hans
; TITLE OF INVENTION: IMMUNOCONJUGATES AND HUMANIZED
; TITLE OF INVENTION: ANTIBODIES SPECIFIC FOR B-CELL LYMPHOMA AND LEUKEMIA CELLS
; NUMBER OF SEQUENCES: 21
; CORRESPONDENCE ADDRESS:
; ADDRESSEE: Foley & Lardner
; STREET: 3000 K Street, N.W., Suite 500
; CITY: Washington, D.C.
; COUNTRY: USA
; ZIP: 20007-5109
; COMPUTER READABLE FORM:
; MEDIUM TYPE: Floppy disk
; COMPUTER: IBM PC compatible
; OPERATING SYSTEM: PC-DOS/MS-DOS
; SOFTWARE: PatentIn Release #1.0, Version #1.25
; CURRENT APPLICATION DATA:
; APPLICATION NUMBER: US/08/690,102A
; FILING DATE: 01-JUL-1996
; CLASSIFICATION: 424
; PRIOR APPLICATION DATA:
; APPLICATION NUMBER: US 08/289,576
; FILING DATE: 12-AUG-1994
; ATTORNEY/AGENT INFORMATION:
; NAME: SAXE, Bernhard D.
; REGISTRATION NUMBER: 28,665
; REFERENCE/DOCKET NUMBER: 18733/463/IMIN
; TELECOMMUNICATION INFORMATION:
; TELEPHONE: (202)672-5300
; TELEFAX: (202)672-5399
; TELEX: 904136
; INFORMATION FOR SEQ ID NO: 4:
; SEQUENCE CHARACTERISTICS:
; LENGTH: 116 amino acids
; TYPE: amino acid
; TOPOLOGY: linear
; MOLECULE TYPE: protein
US-08-690-102A-4

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Query Match          100.0%; Score 620; DB 1; Length 116;
Best Local Similarity 100.0%;
Matches 116; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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Qy      61 NQNFKDKATLTADKSSSTAYMQLSSLTSEDSAVVYCCARRDITTFYWGQGTTTLTVSS 116
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Db      61 NQNFKDKATLTADKSSSTAYMQLSSLTSEDSAVVYCCARRDITTFYWGQGTTTLTVSS 116

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<!--StartFragment-->RESULT 1
US-08-690-102A-2
; Sequence 2, Application US/08690102A
; Patent No. 5789554
; GENERAL INFORMATION:
; APPLICANT: LEUNG, Shui-on
; APPLICANT: HANSEN, Hans
; TITLE OF INVENTION: IMMUNOCONJUGATES AND HUMANIZED
; TITLE OF INVENTION: ANTIBODIES SPECIFIC FOR B-CELL LYMPHOMA AND LEUKEMIA CELLS
; NUMBER OF SEQUENCES: 21
; CORRESPONDENCE ADDRESS:
; ADDRESSEE: Foley & Lardner
; STREET: 3000 K Street, N.W., Suite 500
; CITY: Washington, D.C.
; COUNTRY: USA
; ZIP: 20007-5109
; COMPUTER READABLE FORM:
; MEDIUM TYPE: Floppy disk
; COMPUTER: IBM PC compatible
; OPERATING SYSTEM: PC-DOS/MS-DOS
; SOFTWARE: PatentIn Release #1.0, Version #1.25
; CURRENT APPLICATION DATA:
; APPLICATION NUMBER: US/08/690,102A
; FILING DATE: 01-JUL-1996
; CLASSIFICATION: 424
; PRIOR APPLICATION DATA:
; APPLICATION NUMBER: US 08/289,576
; FILING DATE: 12-AUG-1994
; ATTORNEY/AGENT INFORMATION:
; NAME: SAXE, Bernhard D.
; REGISTRATION NUMBER: 28,665
; REFERENCE/DOCKET NUMBER: 18733/463/IMIN
; TELECOMMUNICATION INFORMATION:
; TELEPHONE: (202)672-5300
; TELEFAX: (202)672-5399
; TELEX: 904136
; INFORMATION FOR SEQ ID NO: 2:
; SEQUENCE CHARACTERISTICS:
; LENGTH: 113 amino acids
; TYPE: amino acid
; TOPOLOGY: linear
; MOLECULE TYPE: protein
US-08-690-102A-2

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Qy      61 ESGVPDRFTGSGSGTDFTLTISRQVEDLAIYYCHQYLVSSWTFGGGKLEIK 112
        ||||||||||||||||||||||||||||||||||||||||||||||||||||
Db      61 ESGVPDRFTGSGSGTDFTLTISRQVEDLAIYYCHQYLVSSWTFGGGKLEIK 112

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<!--StartFragment-->RESULT 1
AAO27198
ID   AAO27198 standard; protein; 123 AA.
XX
AC   AAO27198;
XX
DT   17-SEP-2003 (first entry)
XX
DE   Murine anti-CD22 antibody, RFB4, VH protein.
XX
KW   Framework-patching; complementarity determining region; CDR; mouse;
KW   murine; cytostatic activity; cancer; Non-Hodgkin's lymphoma;
KW   gene therapy; rheumatoid arthritis; FR-patching; RFB4 VH; CD22; antibody.
XX
OS   Mus sp.
XX
FH   Key                Location/Qualifiers
FT   Domain              31..35
FT                          /note= "Complementarity determining region (CDR) 1"
FT   Domain              50..66
FT                          /note= "Complementarity determining region (CDR) 2"
FT   Domain              99..112
FT                          /note= "Complementarity determining region (CDR) 3"
XX
PN   WO2003002607-A1.
XX
PD   09-JAN-2003.
XX
PF   10-JUN-2002; 2002WO-US018512.
XX
PR   27-JUN-2001; 2001US-00892613.
XX
PA   (LEUN/) LEUNG S S.
XX
PI   Leung SS;
XX
DR   WPI; 2003-210245/20.
XX
PT   New re-engineered or framework-patched immunoglobulin, useful for
PT   preparing a composition for treating cancer, preferably Non-Hodgkin's
PT   lymphoma or rheumatoid arthritis.
XX
PS   Example 1; Fig 1a; 66pp; English.
XX
CC   The invention relates to a novel re-engineered or framework (FR)-patched
CC   immunoglobulin, containing the heavy and/or light chain variable region
CC   (VH/VL) sequences from a parent antibody. Within these chains, at least
CC   one of the compartmentalised framework sequences, defined as FR1, FR2,
CC   FR3 and FR4 are replaced, or patched, by the corresponding framework
CC   sequences from the heavy and light chain immunoglobulin region of a
CC   different species. The FR-patched immunoglobulin binds specifically to an
CC   antigen with affinity comparable to, or within 3-fold of, that of the
CC   parent immunoglobulin. The invention discloses the process of FR-patching
CC   which is used to generate re-engineered immunoglobulin chains having one
CC   or more complementarity determining regions (CDR's) from a donor
CC   immunoglobulin and portions of framework sequences from one or more human
CC   or primate immunoglobulins. The molecules obtained demonstrate cytostatic
CC   activity as well as reduced or eliminated immunogenicity, whilst
CC   maintaining the specificity and affinity of the parent antibody. The FR-
CC   patched immunoglobulin is useful during the preparation of a composition
CC   for treating cancer, preferably Non-Hodgkin's lymphoma and also during
CC   the treatment of rheumatoid arthritis. Furthermore, the molecules of the
CC   invention may also prove useful in gene therapy. The current sequence is
CC   that of the murine anti-CD22 antibody, RFB4, VH protein of the invention

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XX
SQ   Sequence 123 AA;

      Query Match          100.0%;   Score 648;   DB 1;   Length 123;
      Best Local Similarity 100.0%;
      Matches 123;   Conservative    0;   Mismatches    0;   Indels    0;   Gaps    0;

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      |||
Db      1  EVQLVESGGGLVKPGGSLKLSCAASGFAFSIYDMSWVRQTPEKRLEWVAYISSGGGITYY 60

Qy      61  PDTVKGGRFTISRDNAKNTLYLQMSSLKSEDTAMYYCARHSGYSSYGVLFAFWGQGT LVT 120
      |||
Db      61  PDTVKGGRFTISRDNAKNTLYLQMSSLKSEDTAMYYCARHSGYSSYGVLFAFWGQGT LVT 120

Qy      121 VSA 123
      |||
Db      121 VSA 123

<!--EndFragment-->

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<!--StartFragment-->RESULT 4

AAO27199

ID AAO27199 standard; protein; 107 AA.

XX

AC AAO27199;

XX

DT 17-SEP-2003 (first entry)

XX

DE Murine anti-CD22 antibody, RFB4, VL protein.

XX

KW Framework-patching; complementarity determining region; CDR; mouse;

KW murine; cytostatic activity; cancer; Non-Hodgkin's lymphoma;

KW gene therapy; rheumatoid arthritis; FR-patching; RFB4 VL; CD22; antibody.

XX

OS Mus sp.

XX

FH Key Location/Qualifiers

FT Domain 24..34

FT /note= "Complementarity determining region (CDR) 1"

FT Domain 50..56

FT /note= "Complementarity determining region (CDR) 2"

FT Domain 89..97

FT /note= "Complementarity determining region (CDR) 3"

XX

PN WO2003002607-A1.

XX

PD 09-JAN-2003.

XX

PF 10-JUN-2002; 2002WO-US018512.

XX

PR 27-JUN-2001; 2001US-00892613.

XX

PA (LEUN/) LEUNG S S.

XX

PI Leung SS;

XX

DR WPI; 2003-210245/20.

XX

PT New re-engineered or framework-patched immunoglobulin, useful for

PT preparing a composition for treating cancer, preferably Non-Hodgkin's

PT lymphoma or rheumatoid arthritis.

XX

PS Example 1; Fig 1b; 66pp; English.

XX

CC The invention relates to a novel re-engineered or framework (FR)-patched
 CC immunoglobulin, containing the heavy and/or light chain variable region
 CC (VH/VL) sequences from a parent antibody. Within these chains, at least
 CC one of the compartmentalised framework sequences, defined as FR1, FR2,
 CC FR3 and FR4 are replaced, or patched, by the corresponding framework
 CC sequences from the heavy and light chain immunoglobulin region of a
 CC different species. The FR-patched immunoglobulin binds specifically to an
 CC antigen with affinity comparable to, or within 3-fold of, that of the
 CC parent immunoglobulin. The invention discloses the process of FR-patching
 CC which is used to generate re-engineered immunoglobulin chains having one
 CC or more complementarity determining regions (CDR's) from a donor
 CC immunoglobulin and portions of framework sequences from one or more human
 CC or primate immunoglobulins. The molecules obtained demonstrate cytostatic
 CC activity as well as reduced or eliminated immunogenicity, whilst
 CC maintaining the specificity and affinity of the parent antibody. The FR-
 CC patched immunoglobulin is useful during the preparation of a composition
 CC for treating cancer, preferably Non-Hodgkin's lymphoma and also during
 CC the treatment of rheumatoid arthritis. Furthermore, the molecules of the
 CC invention may also prove useful in gene therapy. The current sequence is
 CC that of the murine anti-CD22 antibody, RFB4, VL protein of the invention

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XX
SQ   Sequence 107 AA;

      Query Match          99.5%;   Score 559;   DB 1;   Length 107;
      Best Local Similarity 99.1%;
      Matches 106;   Conservative    1;   Mismatches    0;   Indels    0;   Gaps    0;

Qy      1  DIQMTQTSSLSASLGDRVTISCRASQDISNYLNWYQKPDGTVKLLIYYTSLHSGVPS 60
      ||||||||||||||||||||||||||||||||||||||||||||||||||||||||
Db      1  DIQMTQTSSLSASLGDRVTISCRASQDISNYLNWYQKPDGTVKLLIYYTSLHSGVPS 60

Qy      61  KFSGSGSGTDYSLTISNLEQEDFATYFCQQGNTLPWTFGGGKLEIK 107
      :||||||||||||||||||||||||||||||||||||||||||||||||||
Db      61  RFSGSGSGTDYSLTISNLEQEDFATYFCQQGNTLPWTFGGGKLEIK 107
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